<u>CLAIMS</u>

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- 3 1. A recombinant adenovirus comprising a mutation in the E1B-55K gene that encodes a mutated
- 4 E1B-55K protein comprising a single amino acid mutation, said mutation substantially reducing the ability
- of said E1B-55K mutated protein to bind to the tumor suppressor p53.

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A recombinant adenovirus as described in claim 1, wherein said virus is selected from the group consisting of Onyx 051 and Onyx 053.

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10 3. A recombinant adenovirus as described in claim 2 wherein said virus is Onyx 051.

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12 4. A recombinant adenovirus as described in claim 2 wherein said virus is Onyx 053.

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- 14 5. A recombinant adenovirus as described in claim 1, wherein said virus has the further property of
- substantially retaining late functions of said virus.

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- 17 6. An isolated adenoviral E1B-55K protein comprising a single amino acid mutation wherein said
- mutation is selected from the group consisting of amino acids at positions 240 or 260 of said protein.

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7. An isolated adenoviral E1B-55K protein comprising a single amino acid mutation wherein said mutation is at position 240 of said protein.

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8. An isolated adenoviral E1B-55K protein comprising a single amino acid mutation wherein said mutation is at position 260 of said protein.

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- 9. An isolated polynucleotide wherein said polynucleotide comprises mutated adenoviral DNA that
 encodes a E1B-55K protein, said protein comprising a single amino acid mutation which mutation
- substantially reduces the capacity of said protein to bind to the tumor suppressor, p53.

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30 10. An isolated polynucleotide as described in claim 9, wherein said polynucleotide is RNA.

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- 32 11. A method of treating cancer in a patient in need of said treatment, comprising administering to said
- patient a dose of a recombinant adenovirus, said adenovirus comprising a mutation in the E1B-55K gene
- 34 that encodes a mutated E1B-55K protein comprising a single amino acid mutation, said mutation
- substantially reducing the ability of said E1B-55K mutated protein to bind to the tumor suppressor p53,

and allowing sufficient time for said adenovirus to infect said cancer, and repeating said treatment if
 desired.

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4 12. A method as described in claim 11, further comprising administering said recombinant adenovirus

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7 13. A method as described in claim 12, wherein said adenovirus is selected from the group consisting 8 of Onyx 051 or Onyx 053.

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- 10 14. A method of treating cancer in a patient in need of said treatment, comprising administering to said
- patient a dose of an isolated polynucleotide wherein said polynucleotide comprises mutated adenoviral
- 12 DNA that encodes an E1B-55K protein, said protein comprising a single amino acid mutation which
- mutation substantially reduces the capacity of said protein to bind to the tumor suppressor, p53, and
- 14 repeating said treatment if desired.

with a chemotherapeutic.

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16 15. A method of treating cancer as described in claim 14, wherein said polynucleotide is RNA.

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- 18 16. A method of treating cancer as described in claim 15, wherein said polynucleotide encodes said
- 19 E1B-55K protein and said protein comprises a mutation at position 240 of said protein.

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- 21 17. A method of treating cancer as described in claim 15, wherein said polynucleotide encodes said
- 22 E1B-55K protein and said protein comprises a mutation at position 260 of said protein.

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- 24 18. A method as described in claims 16 or 17, further comprising administering said polynucleotide
- 25 with a chemotherapeutic.

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27 19. A method as described in claim 15, wherein said polynucleotide is administered with a liposome.

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